

TABLE 1 50% INHIBITORY CONCENTRATIONS IC ₅₀ FOR FORMULA A			
Example	R ¹	A	mICE IC ₅₀ (μM) CPP32 IC ₅₀ (μM)
4	CH ₃	Ala	>10
7	CH ₃	Pro	>50
10	CH ₃	Val	2.48
13	CH ₃	Leu	5.62
16	CH ₃	Phe	49.8
21	CH ₃	Gly	>50
24	CH ₂ Ph	Ala	>50
27	(CH ₂) ₂ CH=CH ₂	Val	1.45
30	CH ₂ CO ₂ H	Ala	>50
33	(CH ₂) ₂ CO ₂ H	Ala	>50
reference	--	--	47.0

Fig. 1

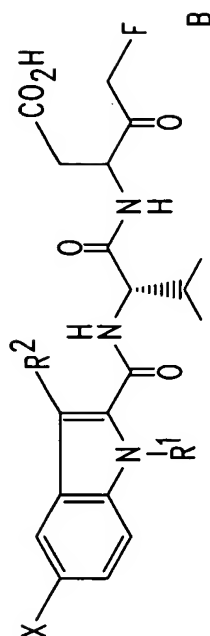


TABLE 2
 DISSOCIATION CONSTANT K_i AND INACTIVATION RATE
 k_3/K_i FOR FORMULA B

Example	R ¹	R ²	X	mICE K_i (μM)	k_3/K_i ($M^{-1}s^{-1}$)	CPP32 K_i (μM)	k_3/K_i ($M^{-1}s^{-1}$)	Mch2 K_i (μM)	k_3/K_i ($M^{-1}s^{-1}$)	Mch5 K_i (μM)	k_3/K_i ($M^{-1}s^{-1}$)
43	CH ₃	CH ₃	H	1.40	2,860	0.960	13,400	0.017	58,800	0.062	21,500
46	CH ₃	Cl	H	1.68	6,150	0.830	25,900	ND	ND	0.099	37,000
49	CH ₃	Cl	F	1.10	7,120	0.493	72,700	0.014	71,400	0.054	52,500
52	(CH ₂) ₃ Ph	H	H	0.133	45,100	0.742	33,700	0.024	41,700	0.077	32,500
55	Ph	H	H	0.843	8,900	0.110	74,200	0.036	55,600	0.043	35,300
58	CH ₂ CO ₂ H	H	H	0.327	16,800	0.125	58,700	0.051	19,600	0.038	127,000
61	CH ₃	H	H	0.240	41,700	0.520	21,200	0.033	30,300	0.026	38,500
62	CH ₃	CH ₃	F	0.397	7,560	0.113	44,200	0.040	25,000	0.102	29,400
63	(CH ₂) ₂ CH=CH ₂	H	H	0.327	18,300	0.125	56,000	0.104	19,200	0.038	131,600
64	CH ₃	H	F	0.234	21,400	0.180	38,900	0.052	38,500	0.063	47,600
65	CH ₃	CH ₂ CH(CH ₃) ₂	H	4.56	1,540	2.28	7,910	0.023	43,500	0.063	31,700
66	CH ₃	(CH ₂) ₂ Ph	H	0.632	14,200	0.505	21,800	0.038	26,300	0.051	39,200
67	CH ₃	H	OCH ₂ Ph	0.739	14,900	0.346	31,800	0.040	25,000	0.062	16,100
reference	--	--	--	0.015	278,000	0.820	14,600	0.594	3,370	0.018	83,300

Fig. 2

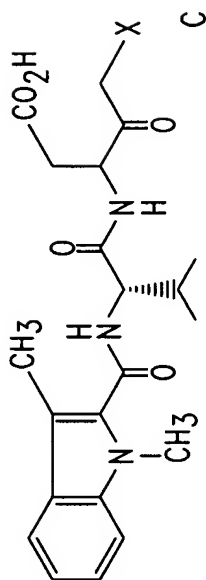


TABLE 3
 DISSOCIATION CONSTANT K_i AND INACTIVATION RATE
 k_3/K_i FOR FORMULA C

Example	X	mICE K_i (μM)	k_3/K_i ($M^{-1}s^{-1}$)	CPP32 K_i (μM)	k_3/K_i ($M^{-1}s^{-1}$)	Mch2 K_i (μM)	k_3/K_i ($M^{-1}s^{-1}$)	Mch5 K_i (μM)	k_3/K_i ($M^{-1}s^{-1}$)
43	F	1.40	2,860	0.960	13,400	0.017	58,800	0.062	21,500
70	OCO (2, 6-di-Cl-C ₆ H ₃)	1.16	3,460	0.052	57,700	0.030	33,300	0.364	2,750
71	OPO (C ₆ H ₅) ₂	0.124	24,200	0.046	65,200	0.060	50,000	0.022	45,500
72	O (1-Ph-3-CF ₃ -pyrazol-5-yl)	0.873	1,150	0.300	16,700	0.050	20,000	1.39	720
73	O 3-CONH ₂ -2-naphthyl)	8.00	250	1.58	0	0.632	1,580	0.213	0
74	O (2-CONH ₂ -1-phenyl)	0.297	3,370	0.419	4,770	0.340	2,940	0.547	0
75	OPO (CH ₃) ₂	4.33	1,850	1.05	7,660	ND	ND	0.663	1,510
reference	--	0.015	278,00	0.820	14,600	0.594	3,370	0.018	83,300

Fig. 3

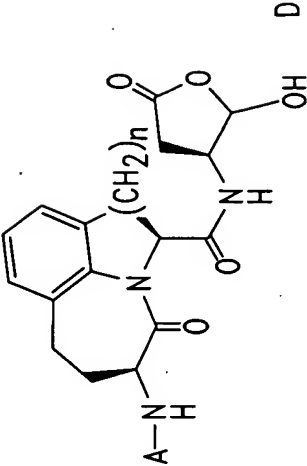
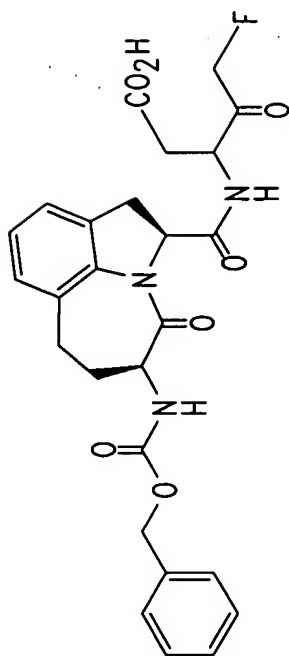


TABLE 4 50% INHIBITORY CONCENTRATIONS IC ₅₀ FOR FORMULA D						
Example No.	A	n	mICE IC ₅₀ (μM)	CPP32 IC ₅₀ (μM)	MCH-2 IC ₅₀ (μM)	MCH-3 IC ₅₀ (μM)
78	Cbz	1	0.019	1.03	40.1	6.96
82	Ac-Asp	1	0.694	0.0014	6.47	0.145
85	succinyl	1	0.571	0.245	1.81	2.83
88	Cbz-Asp	1	0.096	0.00052	ND	0.084
91	dihydrocinnamoyl	1	0.045	0.780	>10	32.6
94	Ac	1	3.07	3.87	>10	>50
100	Benzoyl	1	0.159	8.77	>50	>50
97	1-Naphthoyl	1	0.010	2.91	>50	12.3
103	Cbz	2	0.026	0.437	32.0	1.11
reference	-	-	0.064	47.0	>10	>10

Fig. 4



Example 106

TABLE 5 DISSOCIATION CONSTANT K _i AND INACTIVATION RATE k ₃ /K _i FOR EXAMPLE 106			
Enzyme	Example 106		Reference
	K _i (μM)	k ₃ /K _i (M ⁻¹ s ⁻¹)	K _i (μM) k ₃ /K _i (M ⁻¹ s ⁻¹)
mICE	0.0005	12,000,000	0.015 214,000
CPP32	0.012	960,000	0.820 12,200
MCH-2	0.033	25,000	0.594 2,950
MCH-5	0.022	98,000	0.018 83,300

Fig. 5

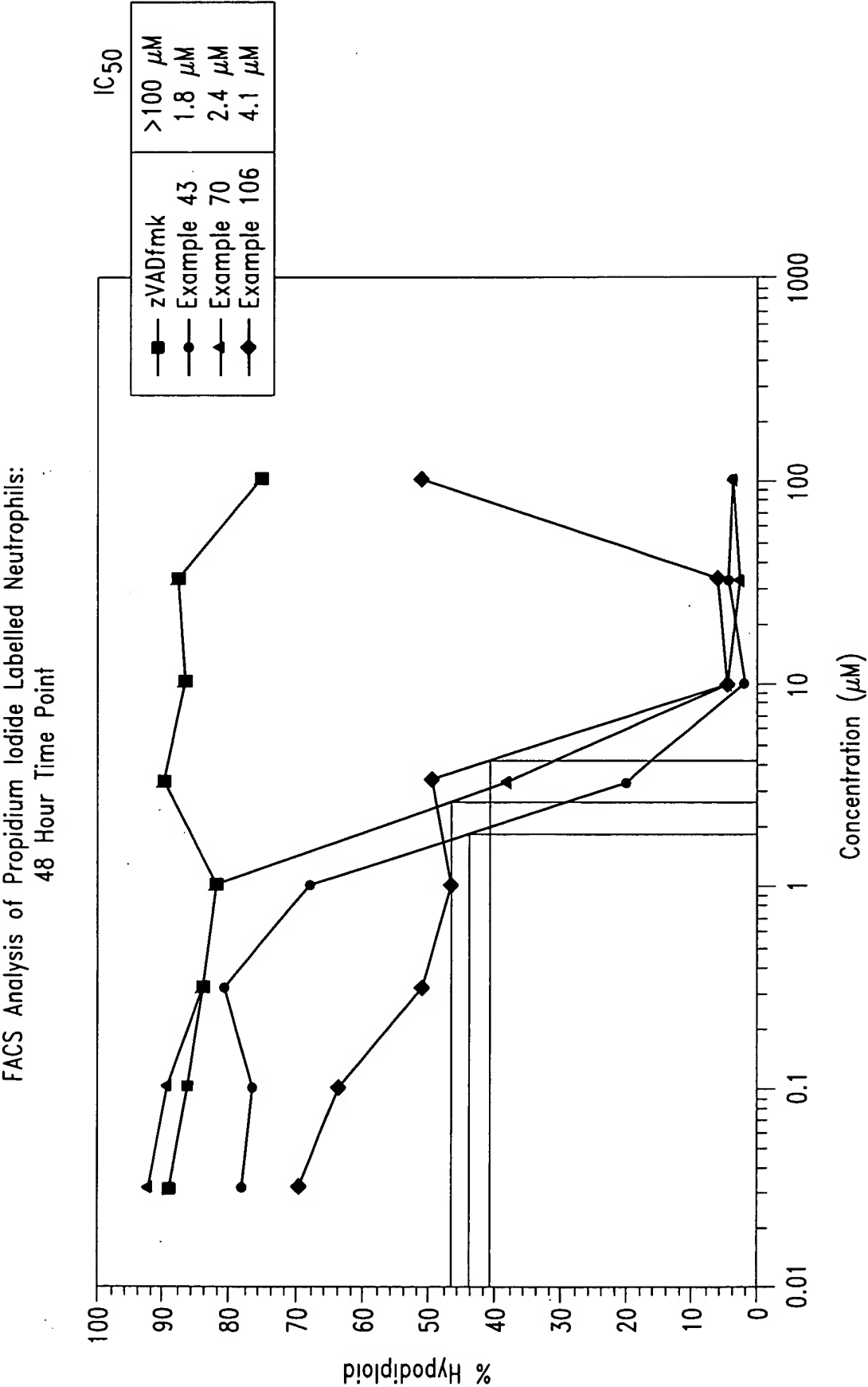


Fig. 6

Neutrophil Survival and Burst Assay

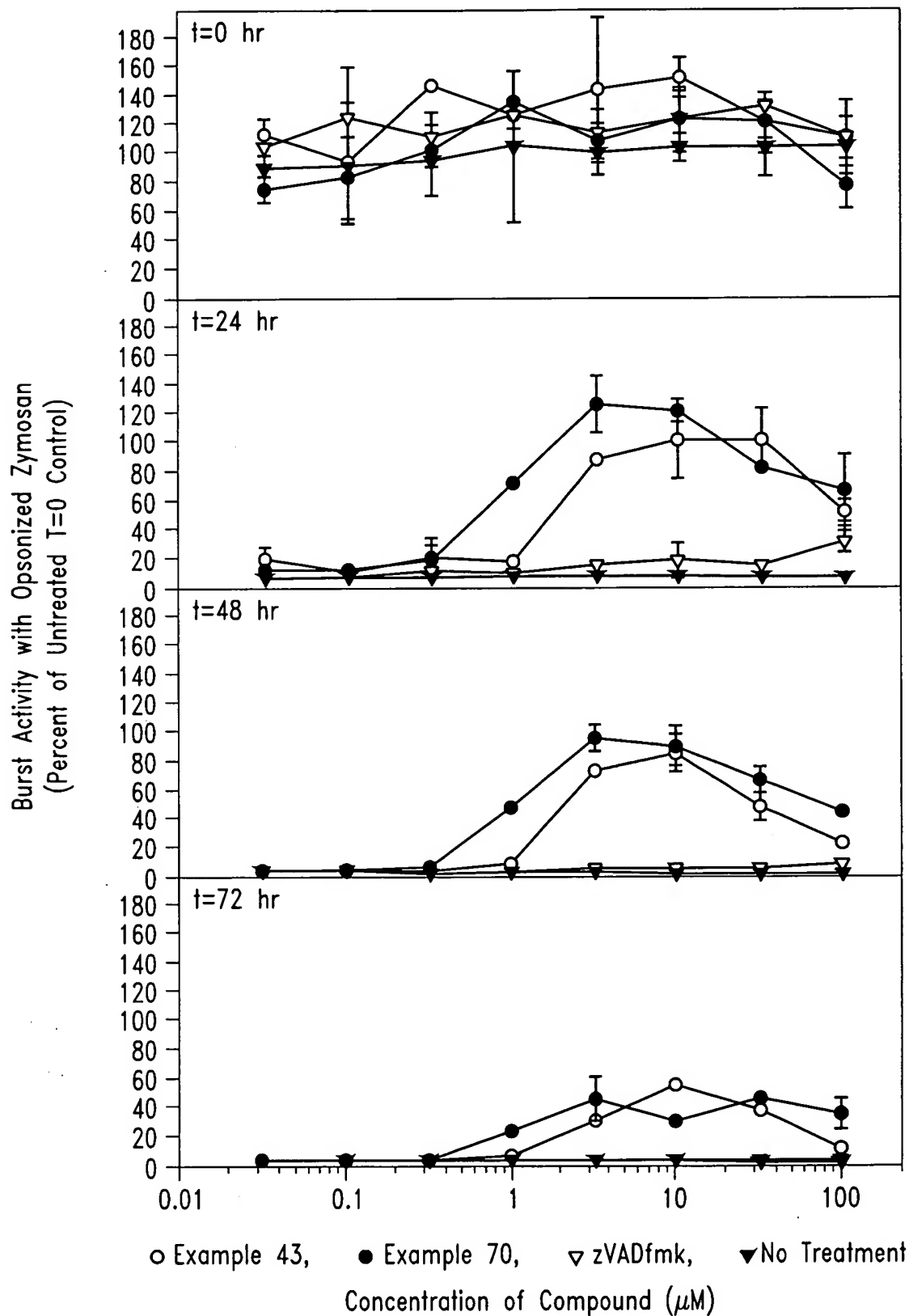


Fig. 7